

AMENDMENTS TO THE SPECIFICATION

Please amend the specification pursuant to 37 C.F.R. 1.121 as follows:

Please replace the paragraph starting on page 8, line 27 with the following paragraph:

-- Preferably, the signal comprises a localisation signal that directs a nascent peptide to the endoplasmic reticulum (ER). A particularly suitable signal is the inclusion of the amino acid sequence Lys-Asp-Glu-Leu (KDEL) (SEQ ID NO.: 1) at the C-terminus of the peptide or polypeptide. Proteins that reside in the lumen of the endoplasmic reticulum (ER), the first compartment for newly made membrane-bound proteins or secreted proteins, are known to possess this short sequence (Munro and Pelham, 1987). If this sequence is deleted or extended by addition of further amino acids, the protein is secreted from the cell rather than retained. --

Please replace the paragraph beginning on page 9, line 9 with the following paragraph:

-- According to a fifth aspect of the invention there is provided a polypeptide comprising a binding region capable of binding to a cell adhesion molecule and a signalling region for subcellular targeting of the polypeptide. Preferably, the polypeptide comprises an antibody or antibody fragment, most preferably a single chain antibody fragment (sFv). The signalling region of choice is a localisation signal for the endoplasmic reticulum. Most preferably, signalling region comprises the amino acid sequence KDEL (SEQ ID NO.: 1) at the C terminus of the polypeptide. --

Please replace the paragraph on page 15, line 18 with the following paragraph:

-- The sFv was amplified from the phagemid vector by PCR (30 cycles, annealing temperature 55°C, 1.5µM Mg²⁺) using the primers:

(5')CAGTCTATGCGGCCCCATTCA(3') (SEQ ID NO.: 2); and

(5')TCCACAGGCGCGCACTCCCAGCCGGGCATGGCCCAGGT(3') (SEQ ID NO.: 3). --

Please replace the paragraph starting on page 17, line 1 with the following paragraph:

-- In transient transfection assays, the cells were assayed 24-72 hours after the start of transfection. The transfected cells were harvested and stained with the monoclonal antibody 10.2C7 specific for porcine VCAM (Celltech, UK) and a second layer reagent labeled with Texas red. In order to identify cells which had taken up the plasmid DNA, the sFv constructs were co-transfected with the vector pEF/GFP/ER which contains a 716bp fragment from p α GFP (Cramer *et al.*, 1996) fused to the KDEL (SEQ. ID NO.: 1) ER-retention signal in the same vector backbone as the sFv constructs. --

Please enter the abstract annexed to this response at Exhibit Tab A.

* * * * *